

Methotrexate entrained in the hyaluronan meshwork

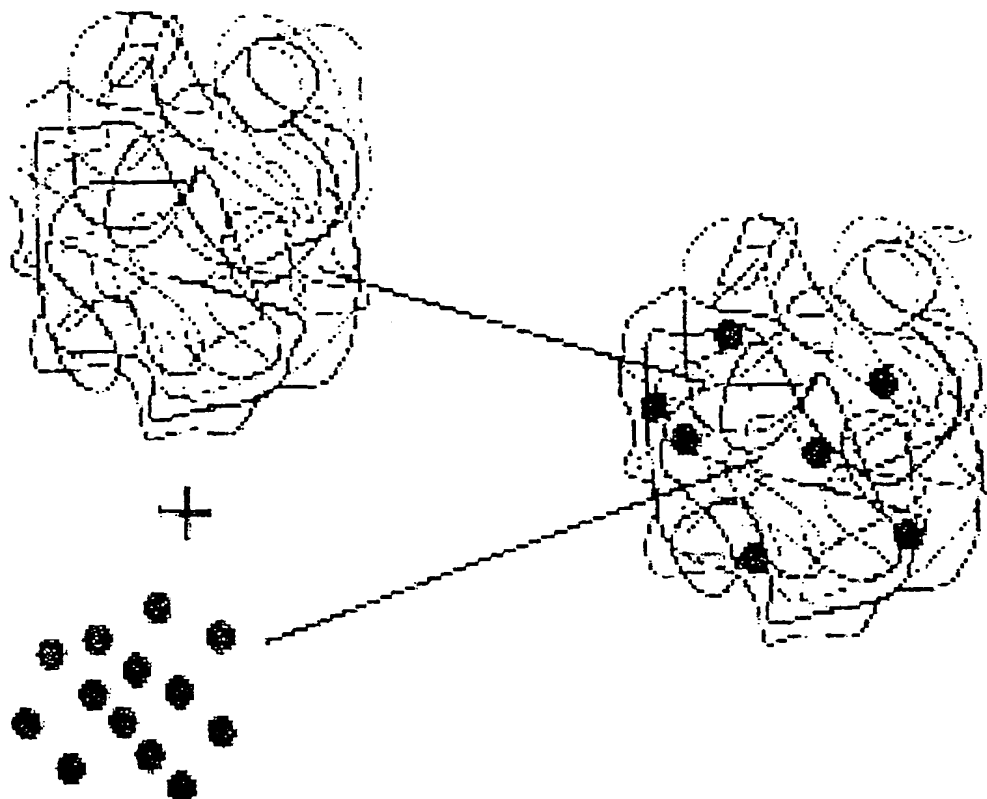
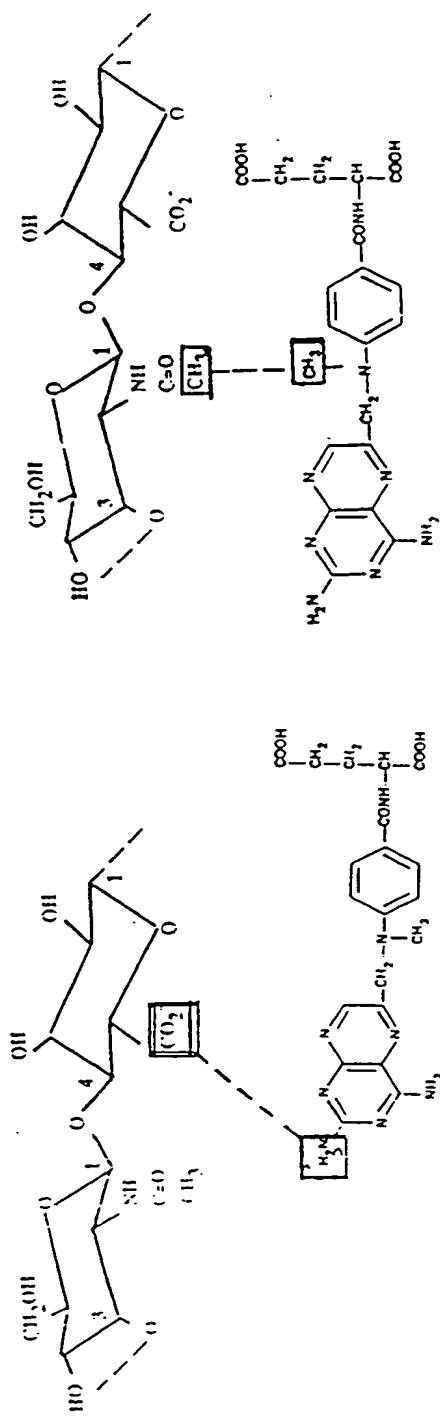


FIGURE 1

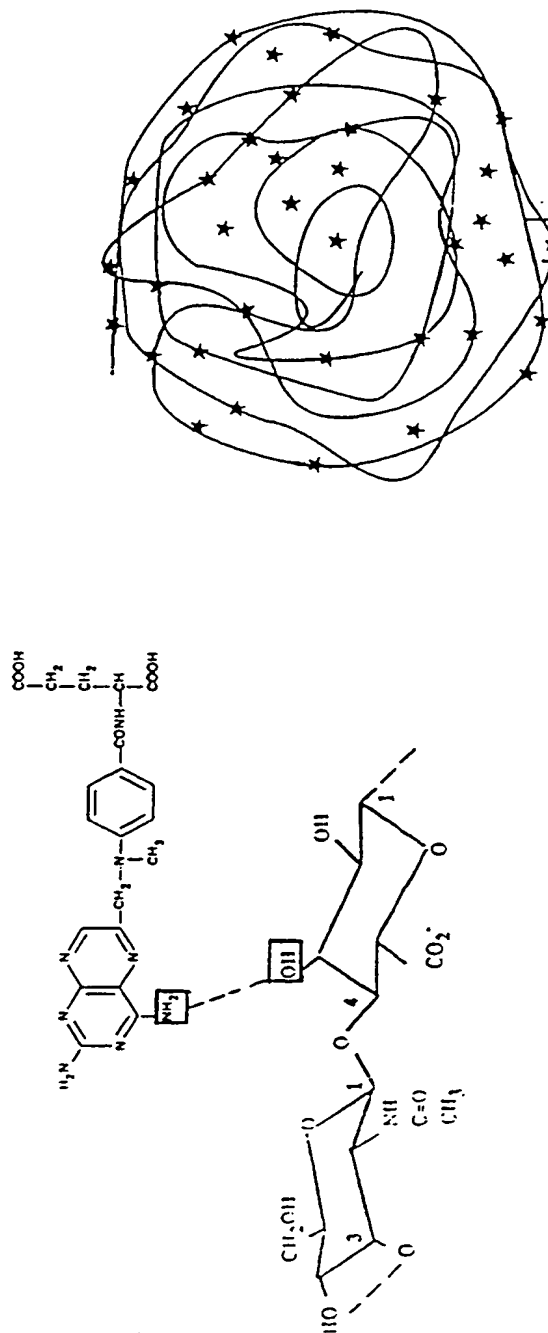
2/28

A: Possible Interactions between Methotrexate and Hyaluronan
i. Ionic bonding
ii. Hydrogen bonding
iii. Hydrophobic bonding



ii. Hydrogen Bonding

B: Entanglement of Methotrexate in Hyaluronan Molecule



..FIGURE . 2.

3/28

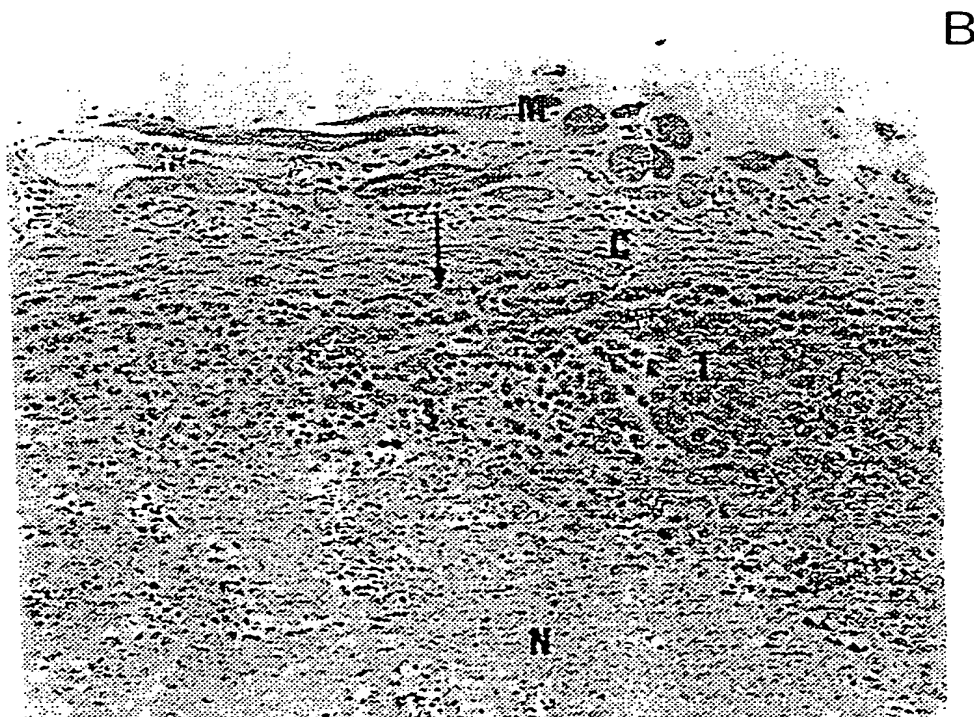
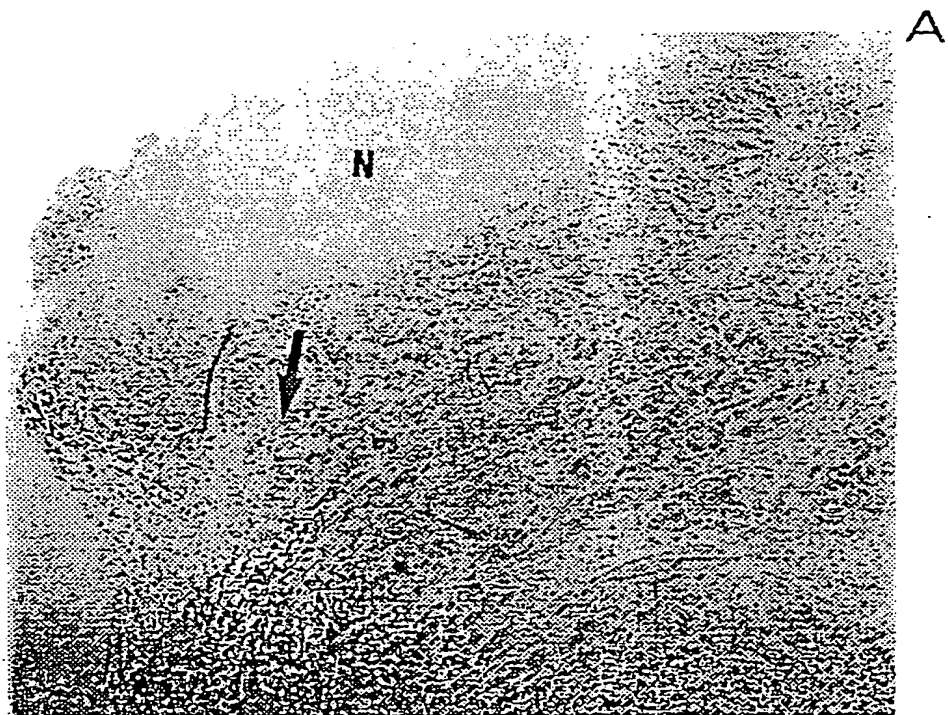


FIGURE 3
SUBSTITUTE SHEET (RULE 26) (RO/AU)

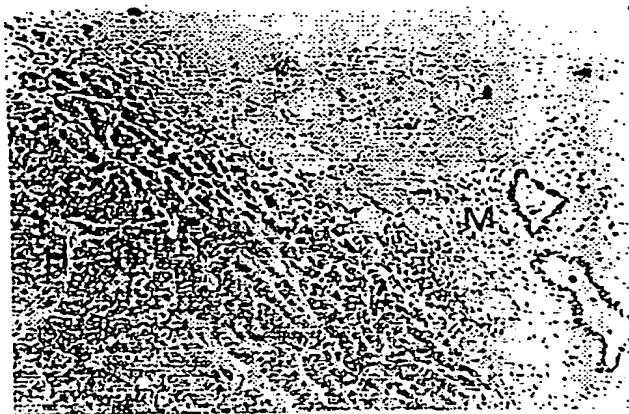
09889203 034302
200700 80268850



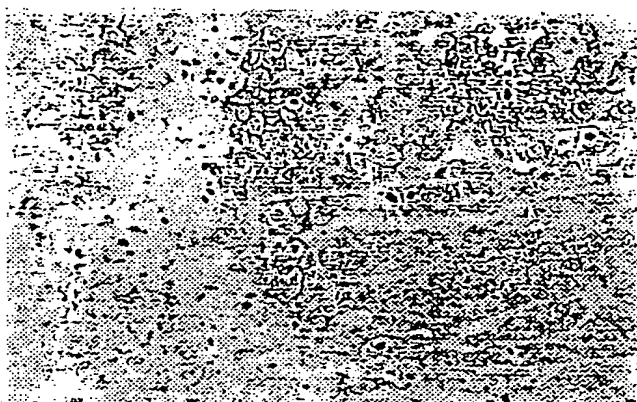
FIGURE 4
SUBSTITUTE SHEET (RULE 26) (RO/AU)

5/28

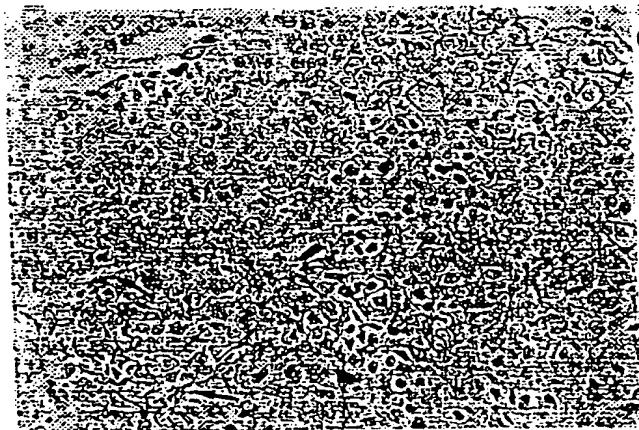
A



B



C



D



FIGURE 5

SUBSTITUTE SHEET (RULE 26) (RO/AU)

6/28

Molecular weight characterisation of hyaluronan
used as a delivery vehicle for methotrexate

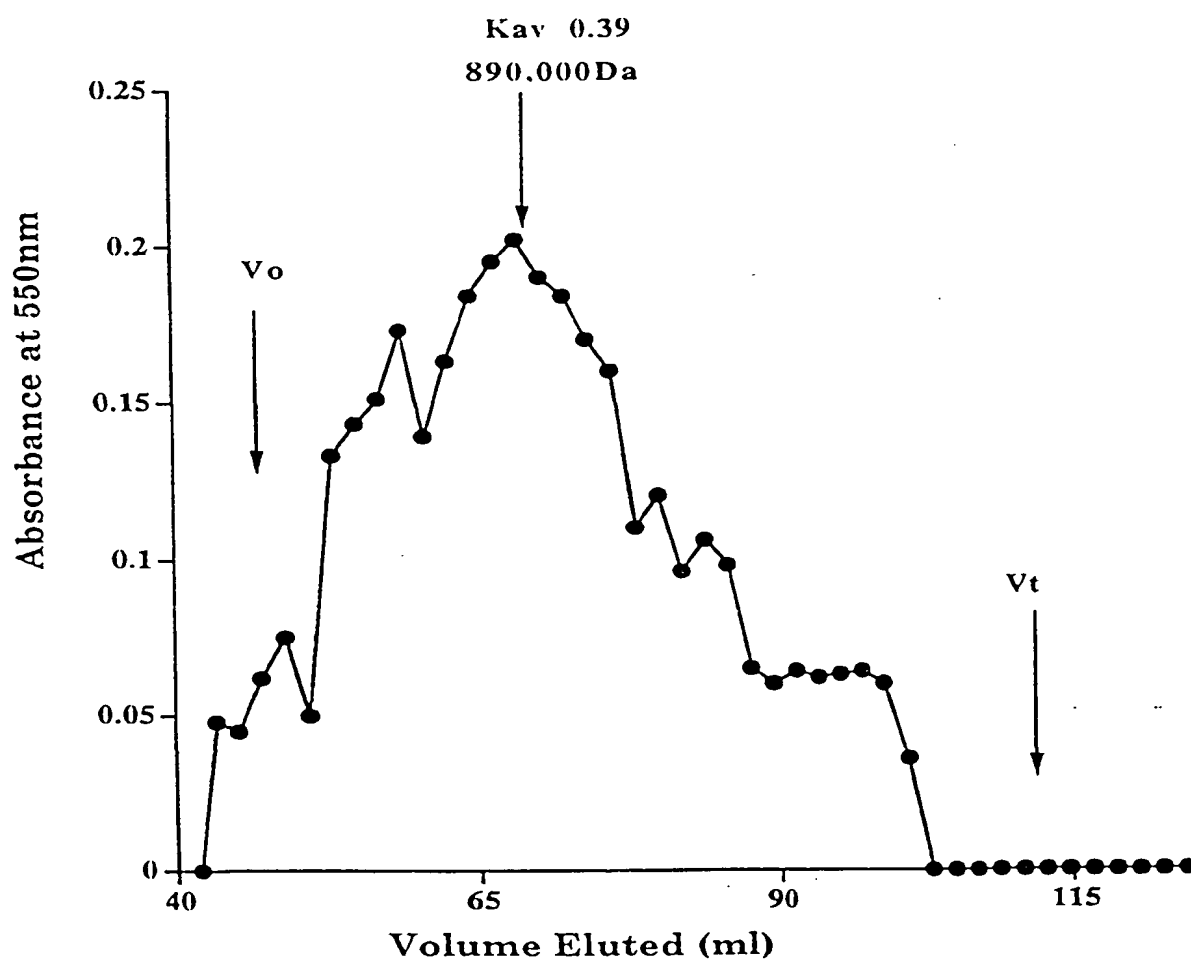


FIGURE 6

7/28

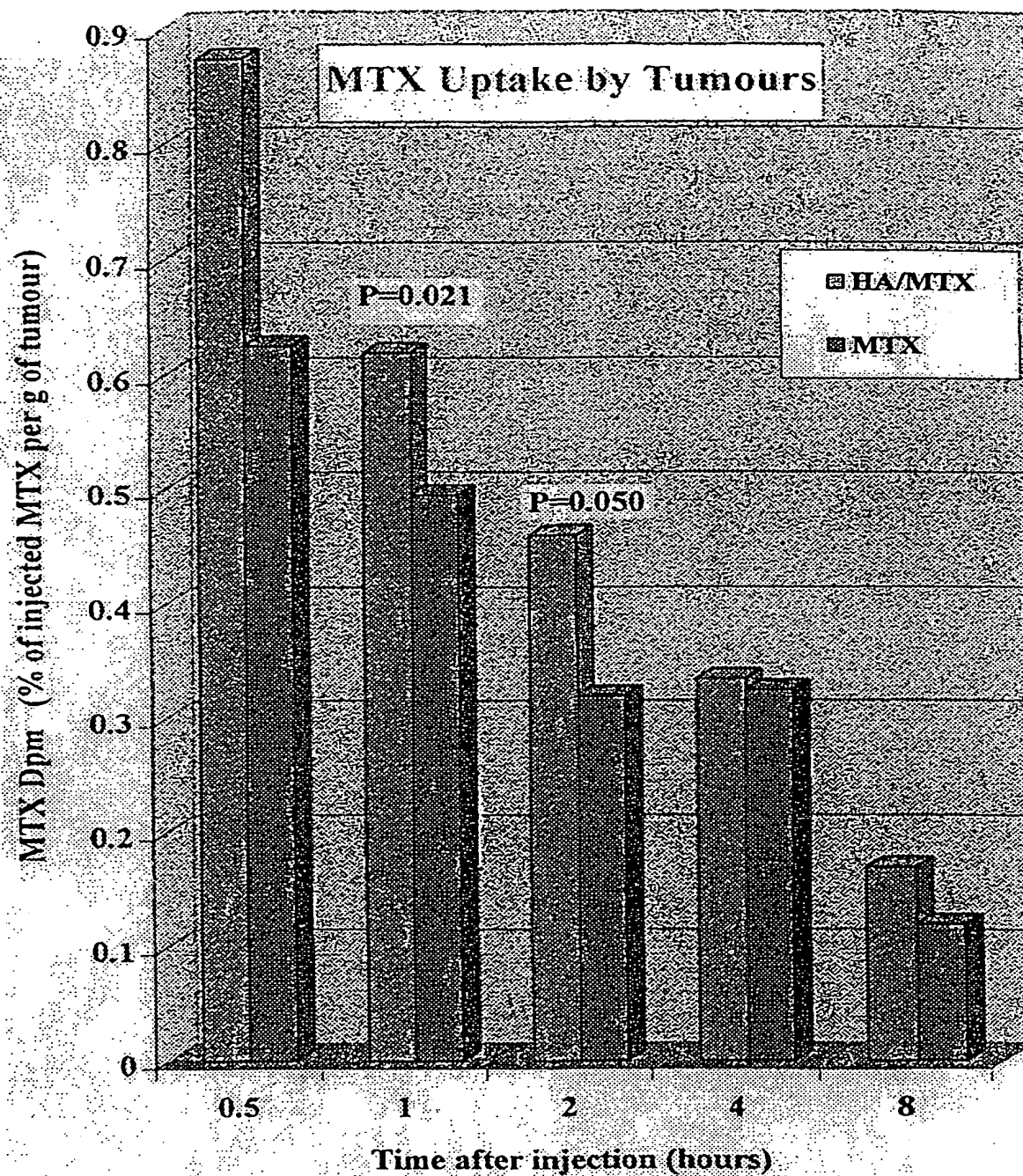


FIGURE 7

SUBSTITUTE SHEET (RULE 26) (RO/AU)

8/28

Increased methotrexate uptake by the liver when drug is
co-administered with hyaluronan

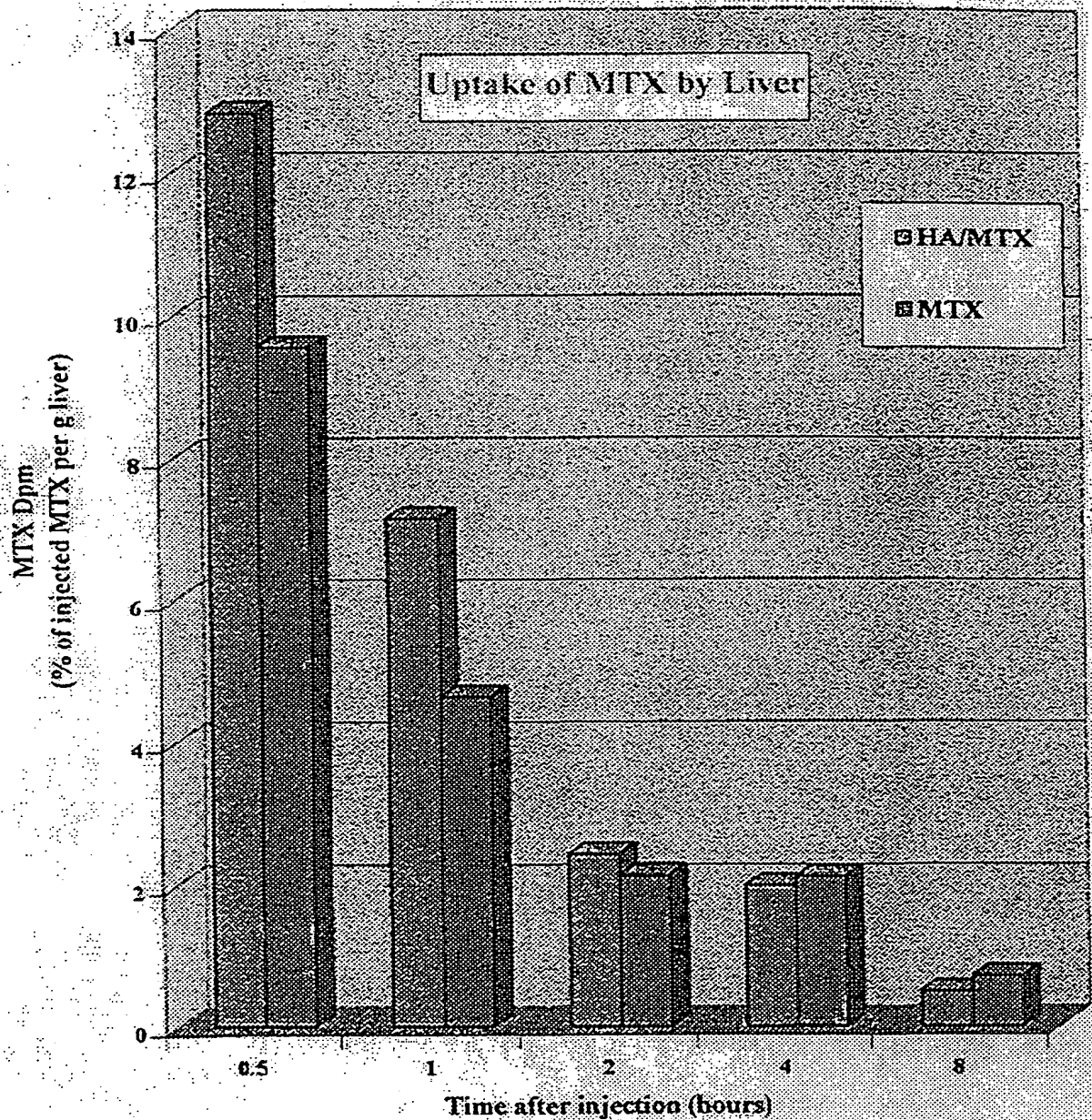


FIGURE 8

9/28

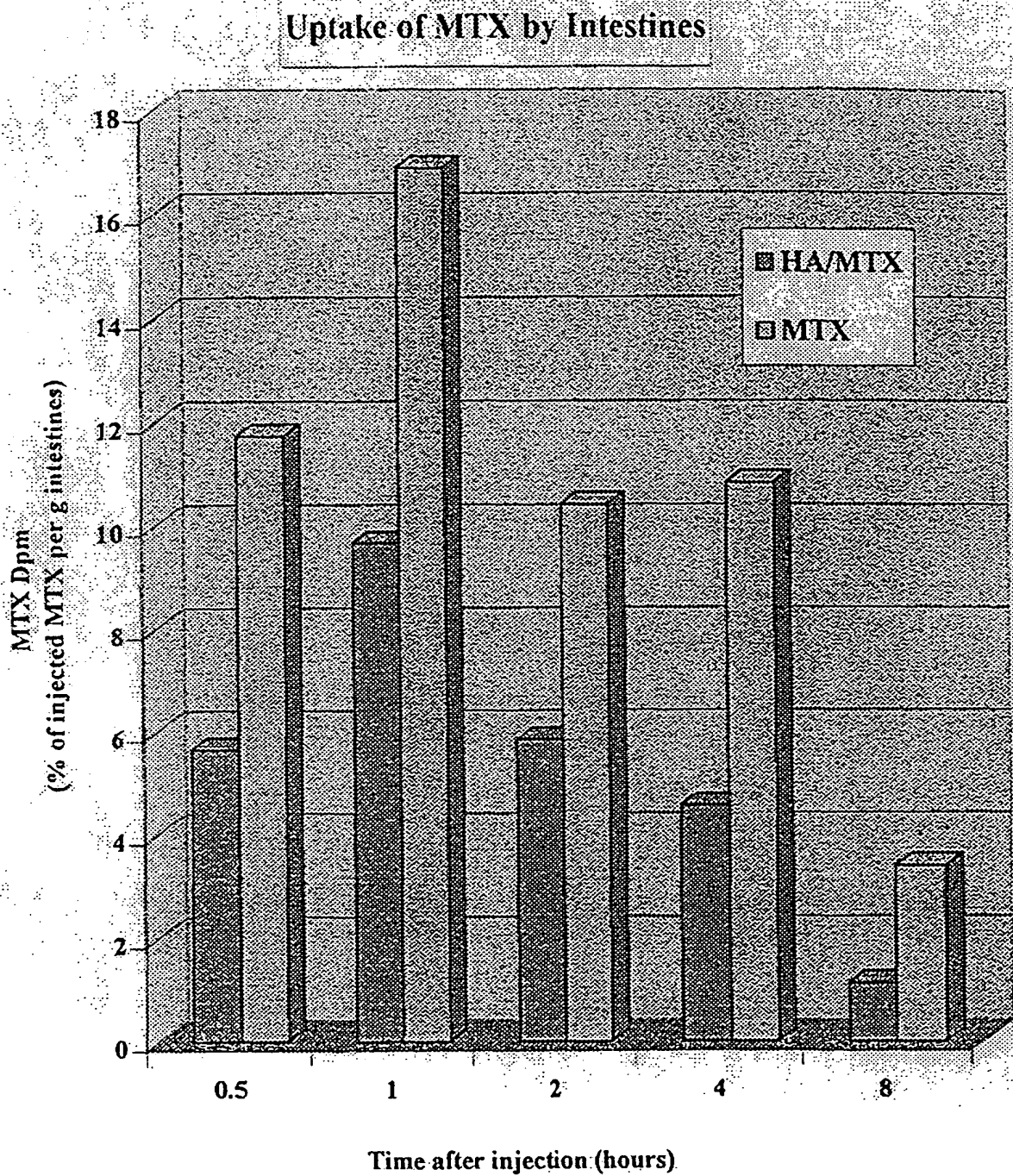


FIGURE 9

10/28

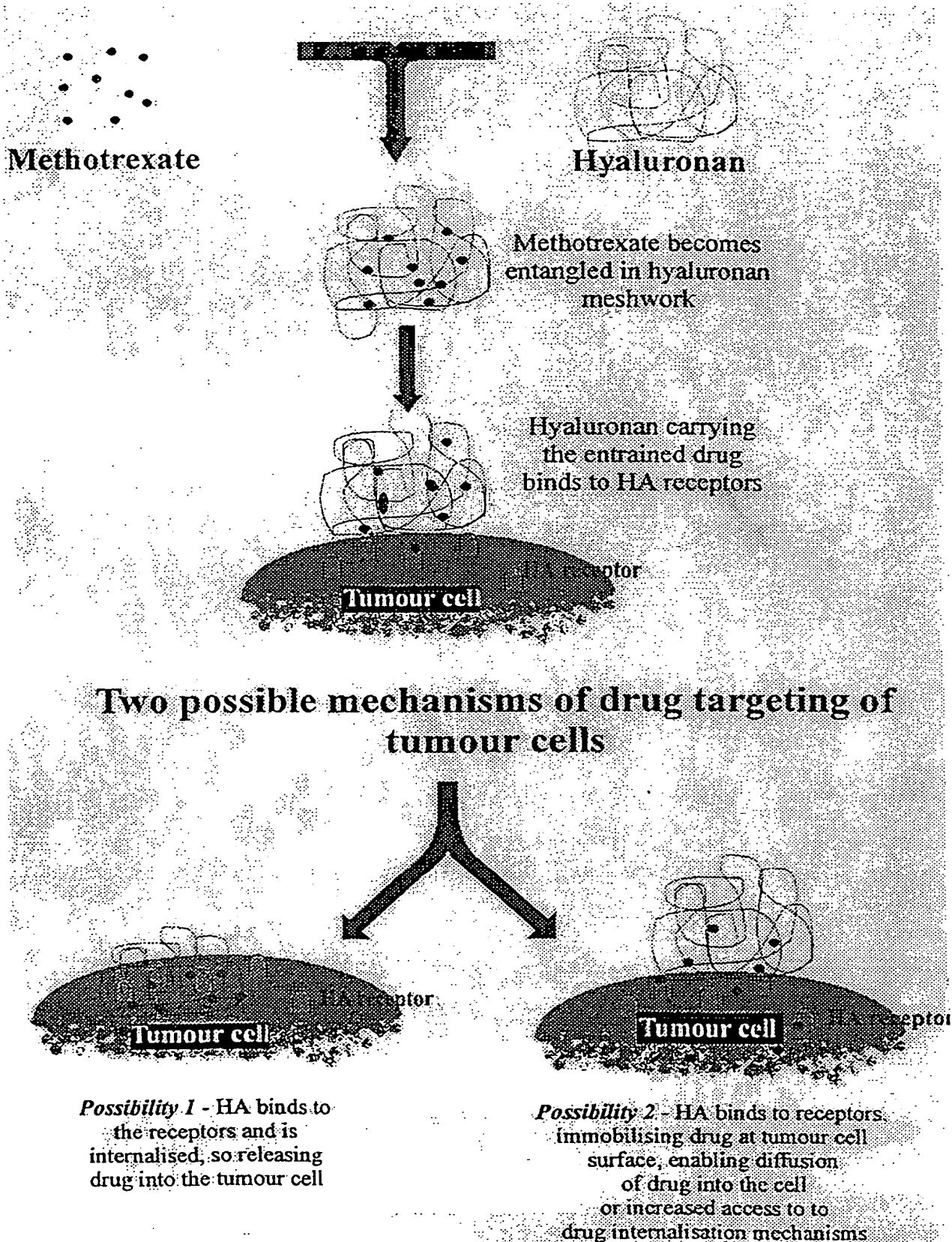
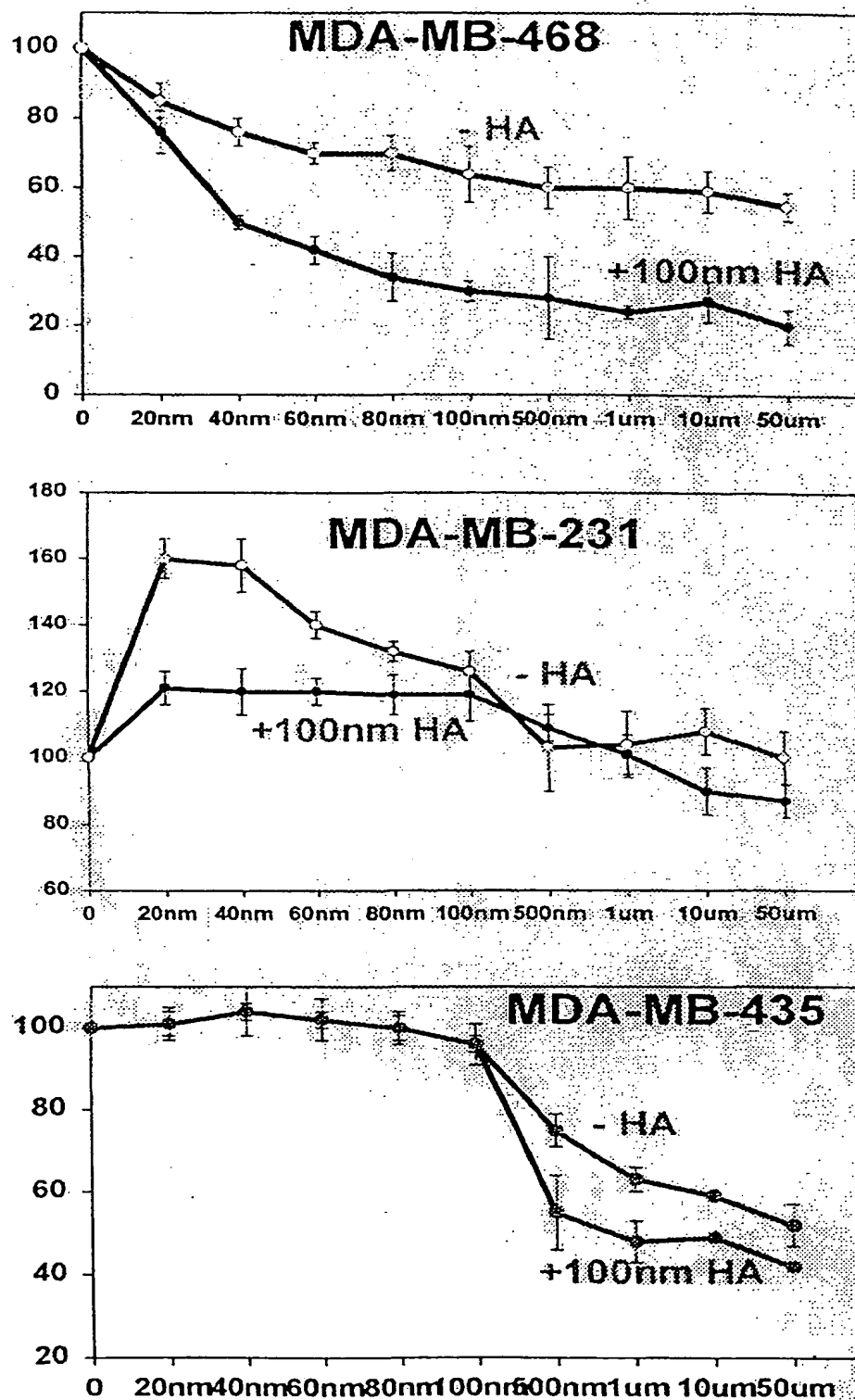


FIGURE 10.

11/28

Percentage of Untreated Control

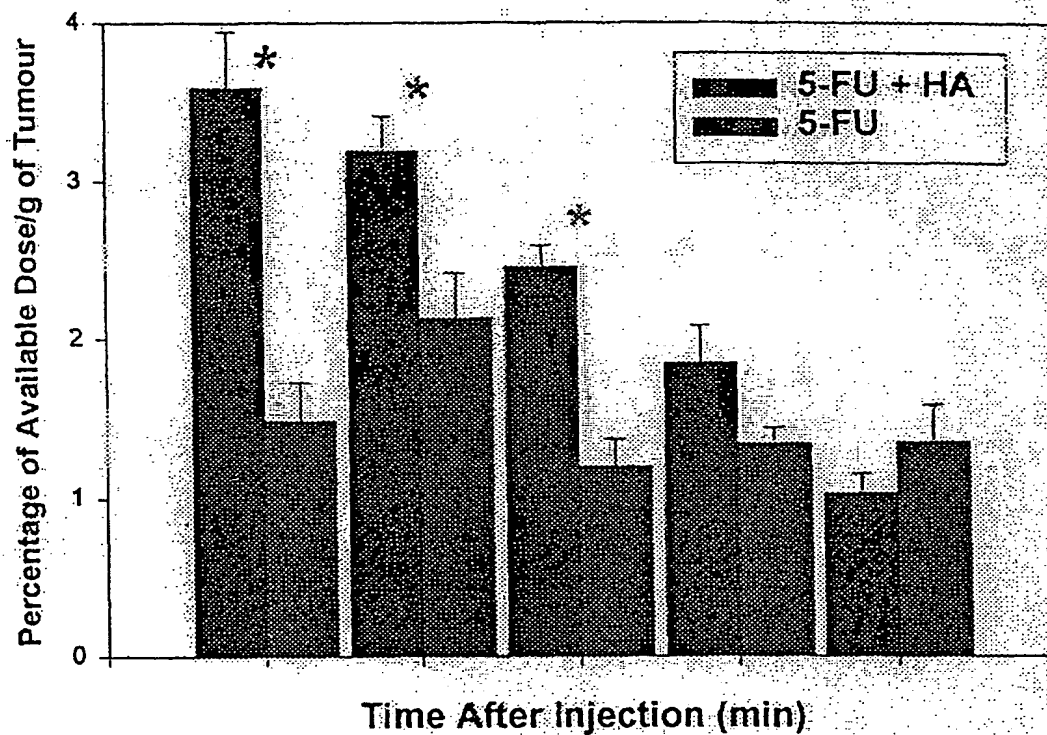


5-FU Concentration

FIGURE 11

12/28

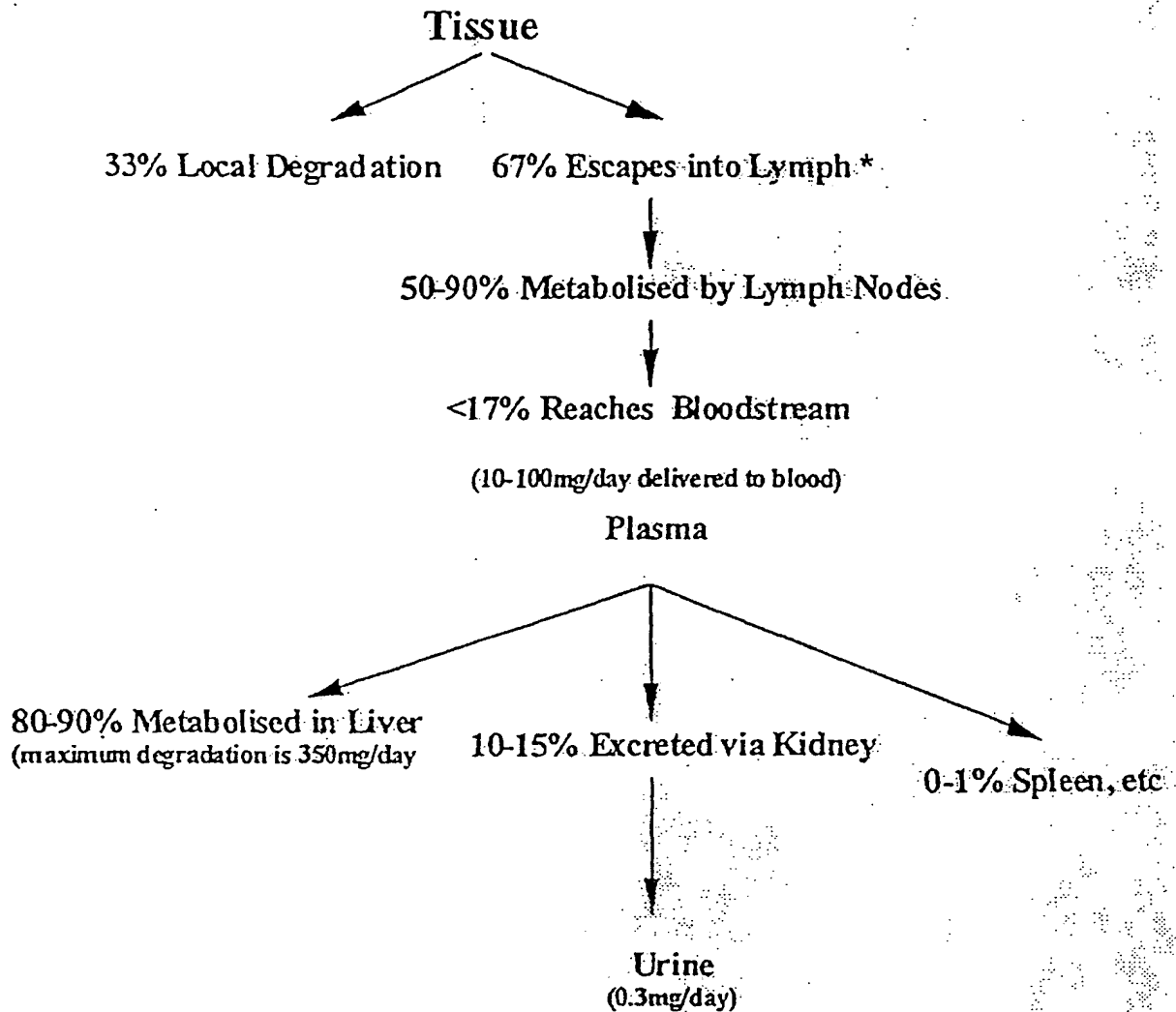
Tumour Uptake of 5-Fluorouracil



* $P = \text{or} < 0.001$, Students t-test

FIGURE 12

13/28



* Local catabolism will almost certainly dominate in very dense tissues, or those without lymphatic drainage; for example, bone and cartilage

FIGURE 13

14/28

Uptake of 5-FU (% of available dose/g tissue)

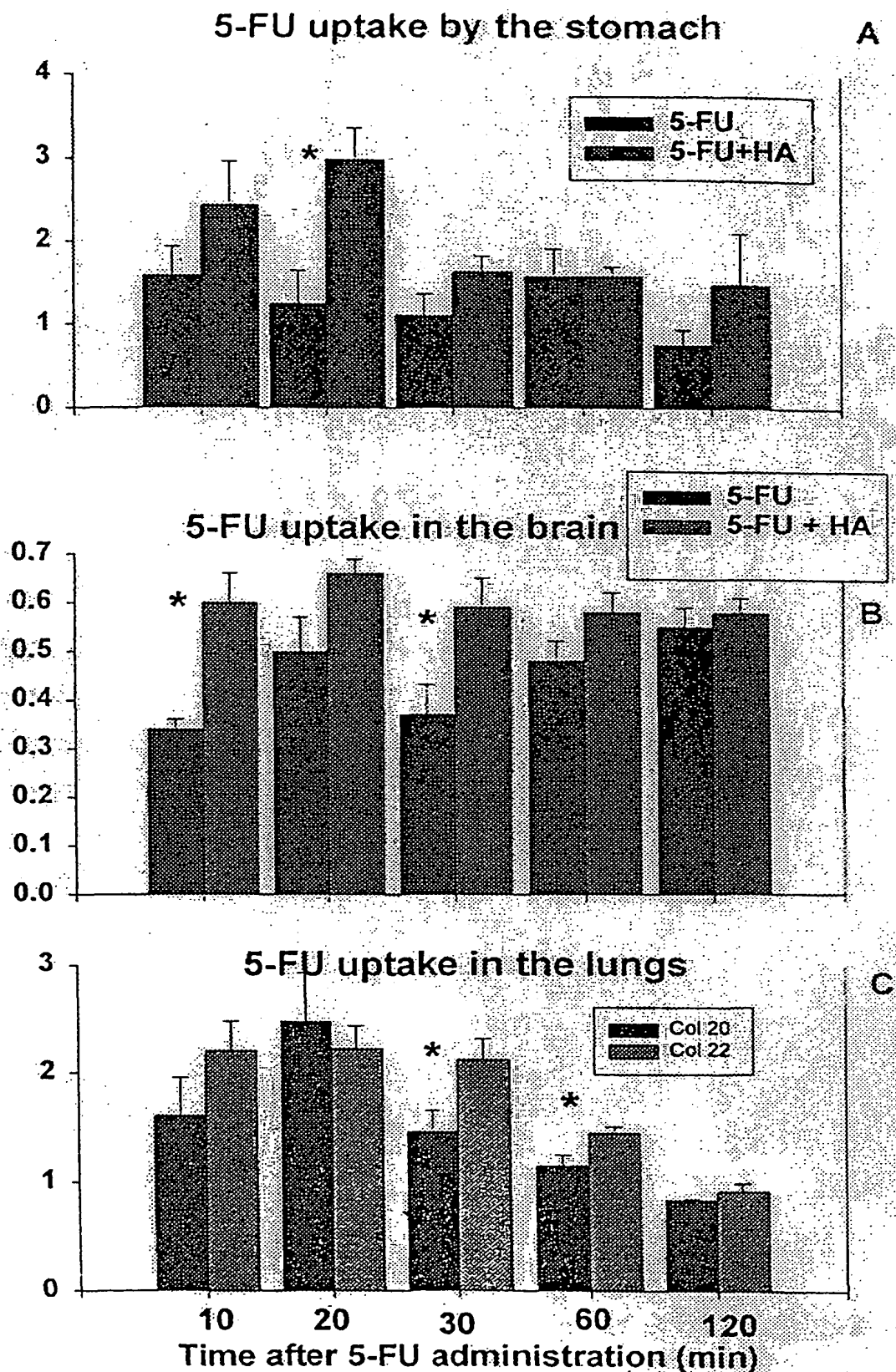


FIGURE 14

15/28

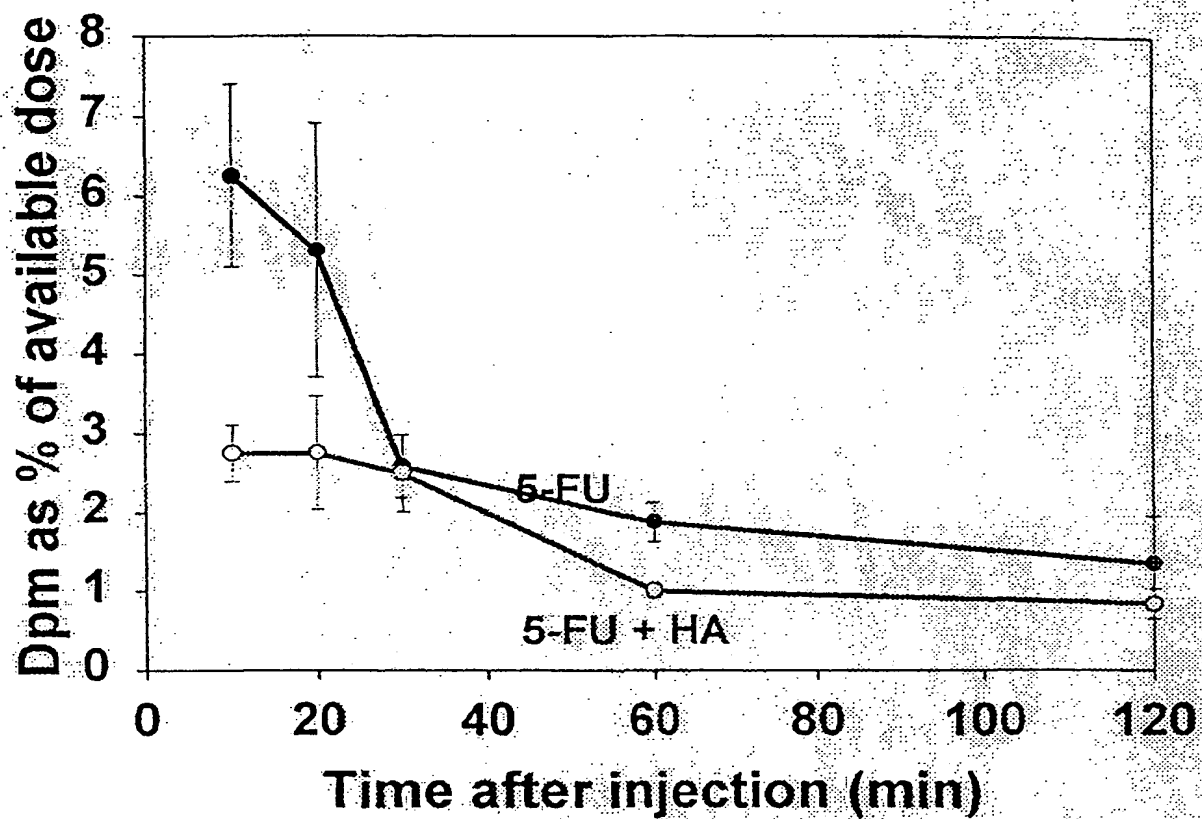


FIGURE 15

16/28

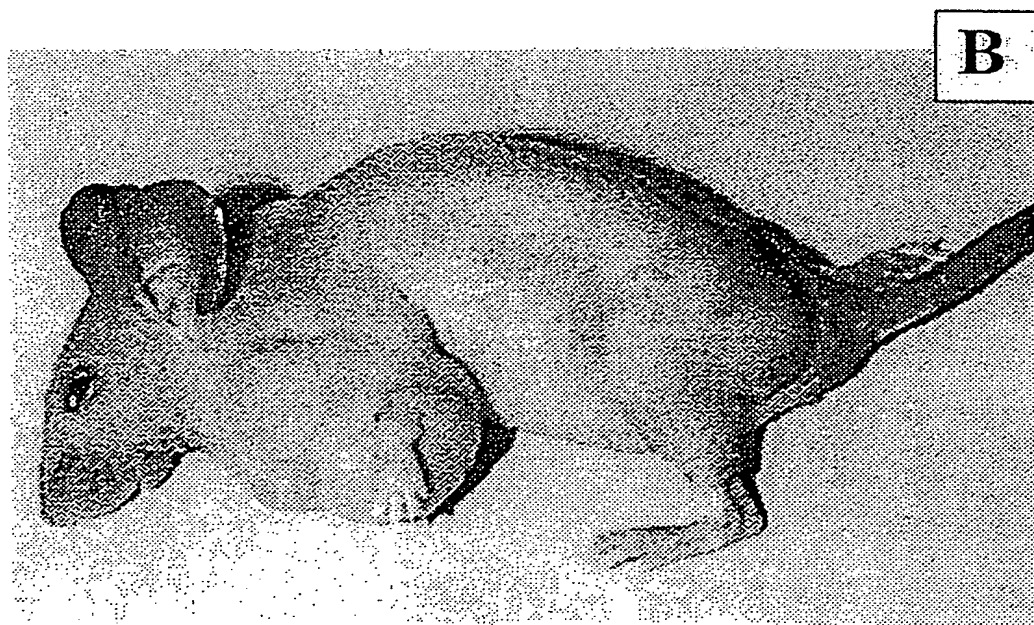
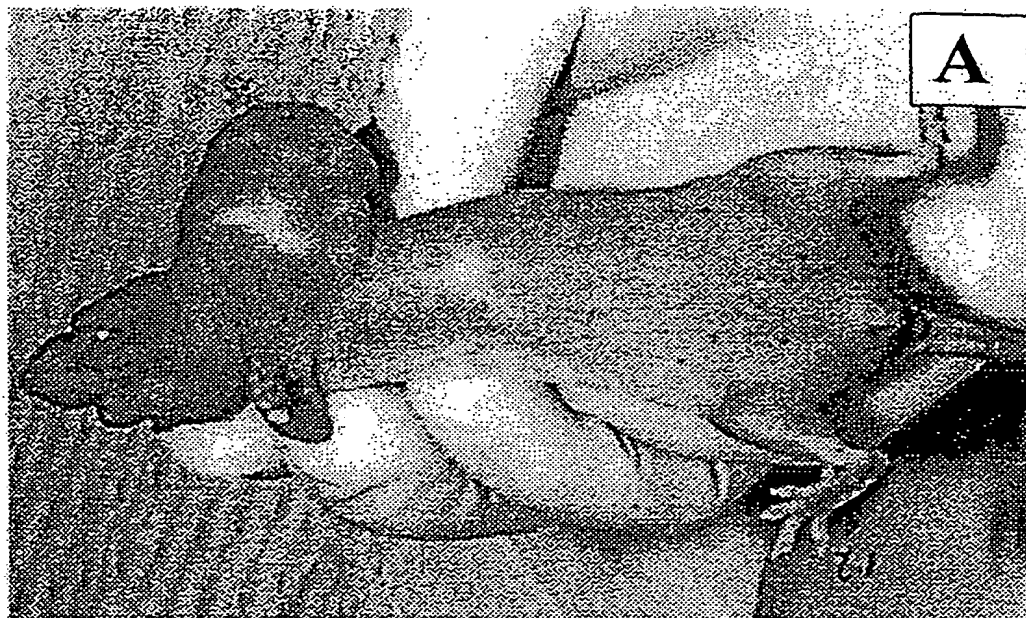


FIGURE 16

17/28

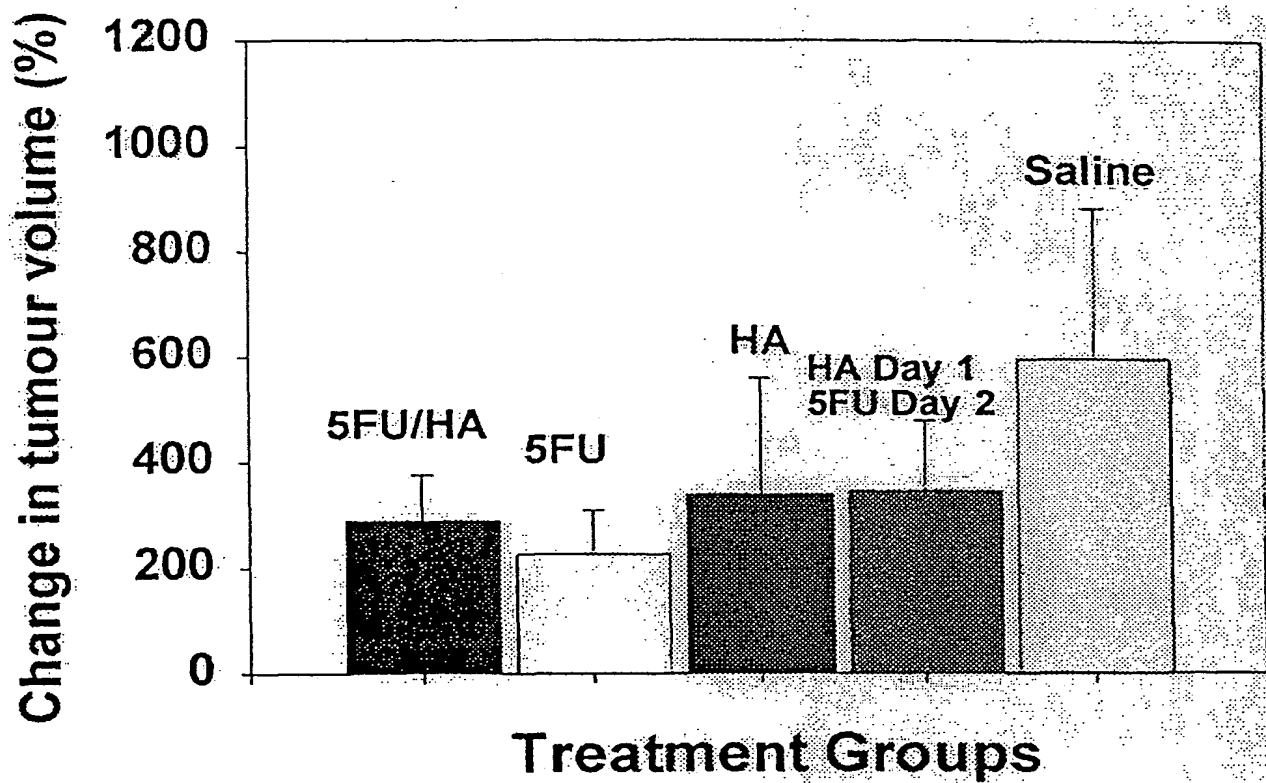


FIGURE 17

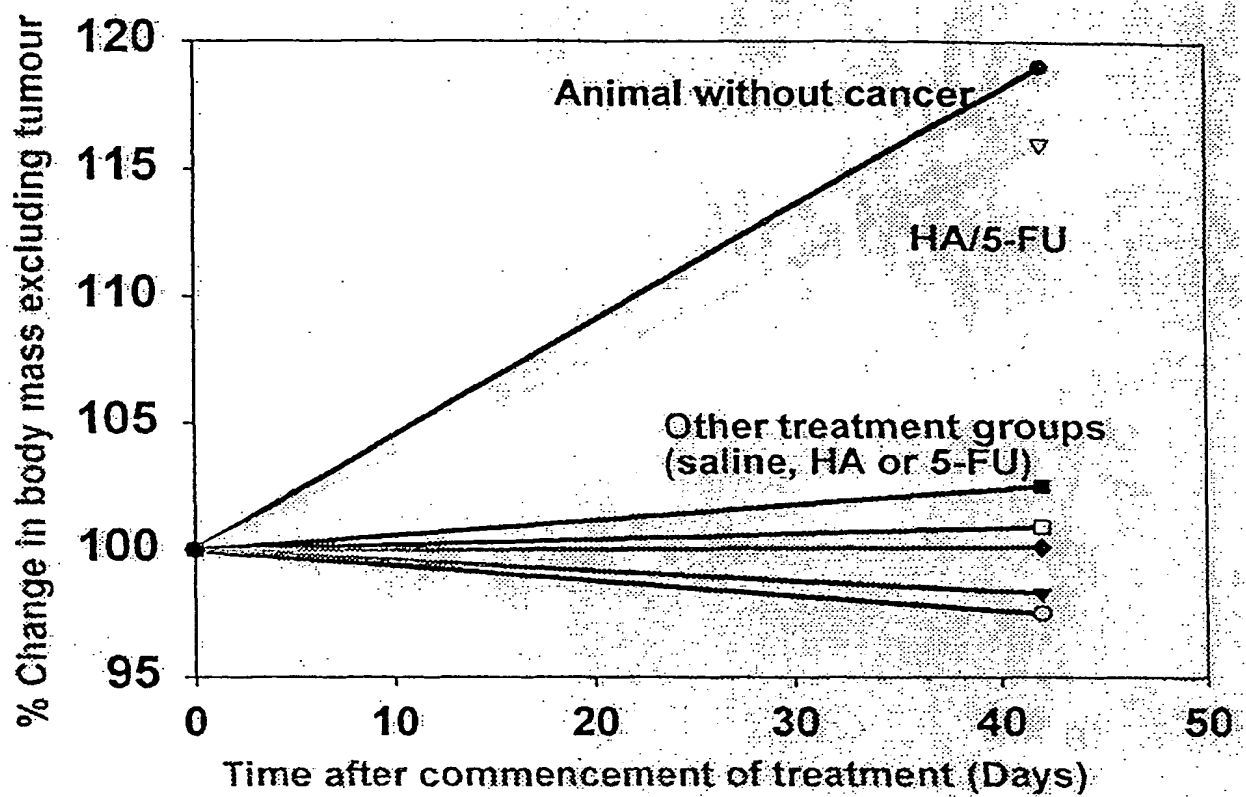


FIGURE 18

19/28

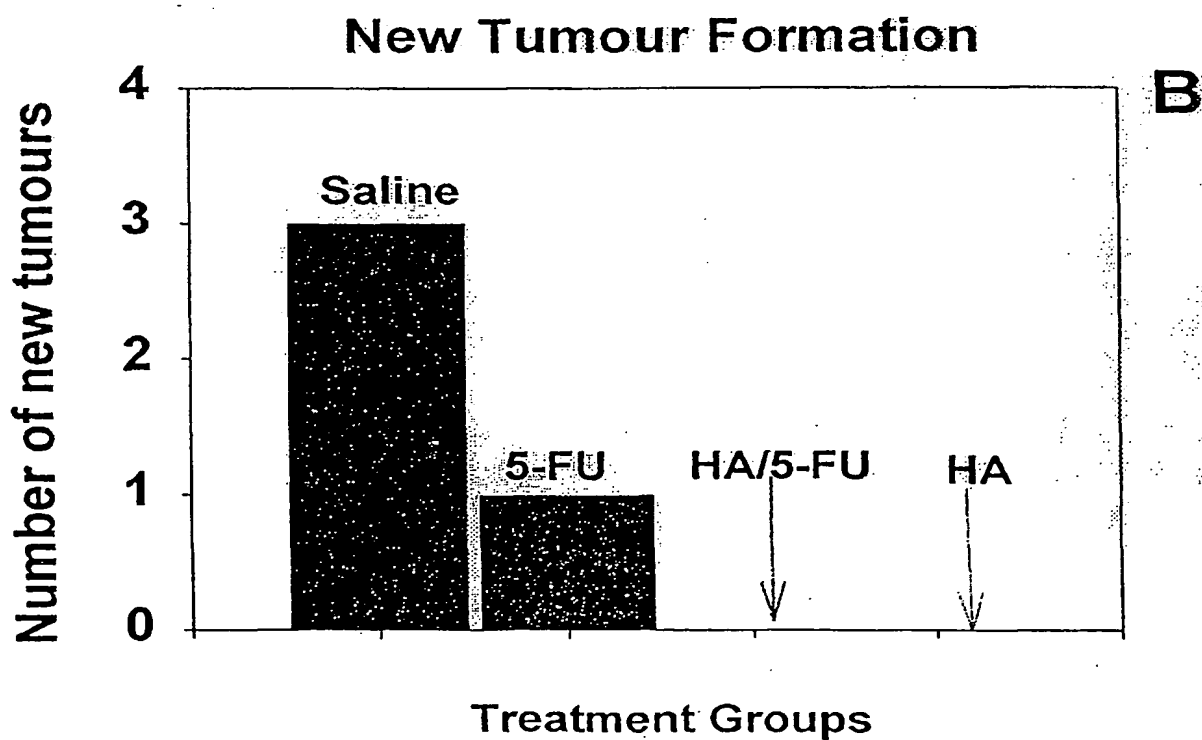
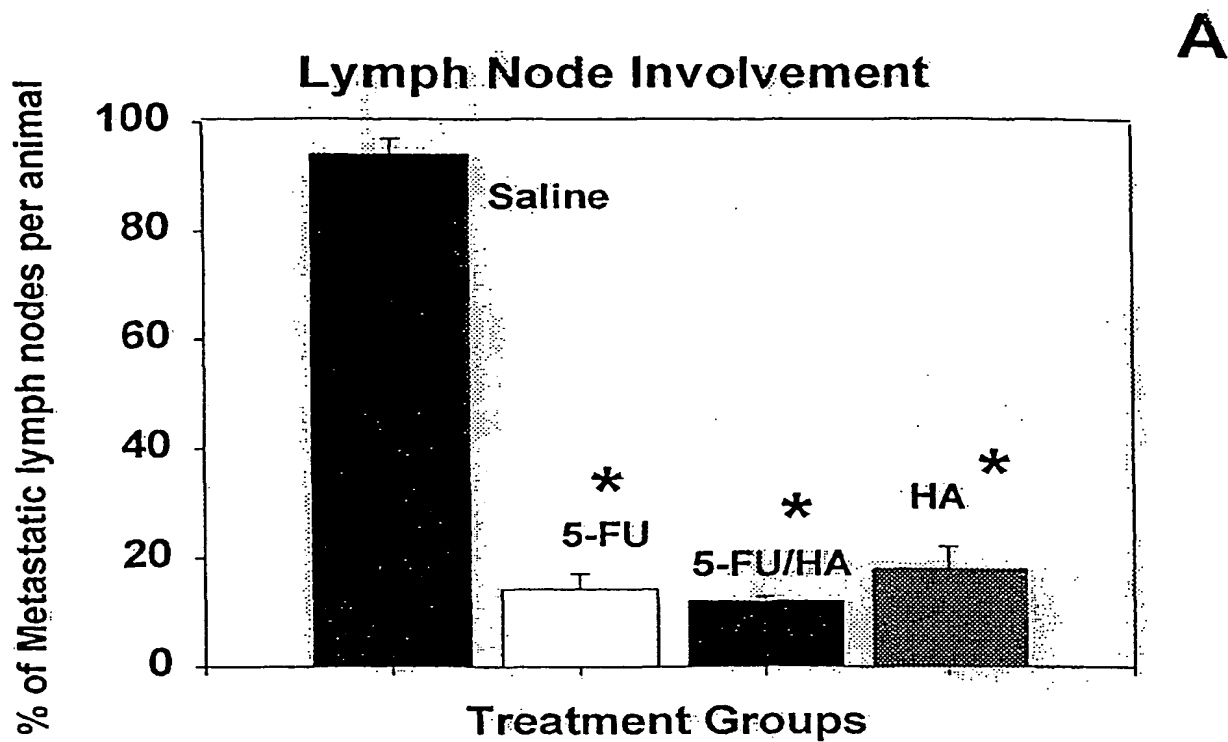


FIGURE 19

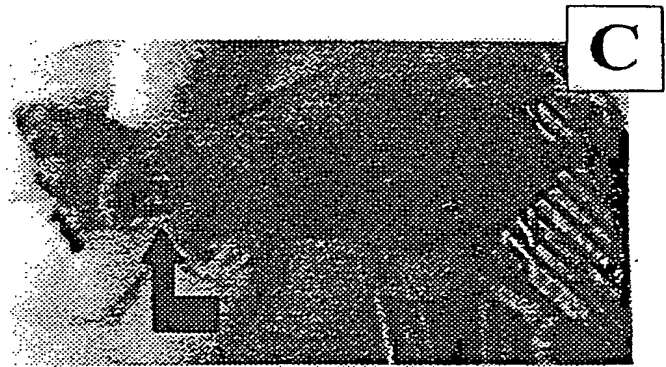
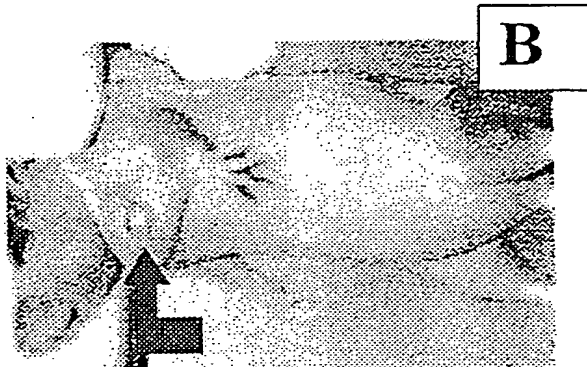
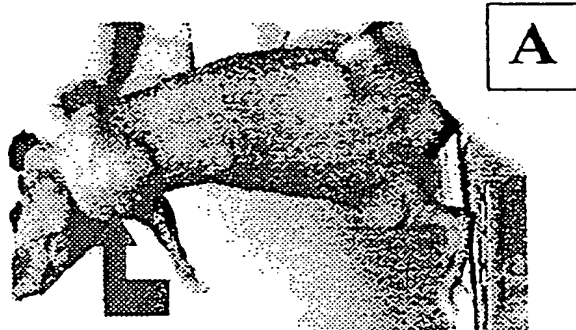


FIGURE 20
SUBSTITUTE SHEET (RULE 26) (RO/AU)

09889203-031302
"FILED" 02268860

6 Month Study: Survival Curves

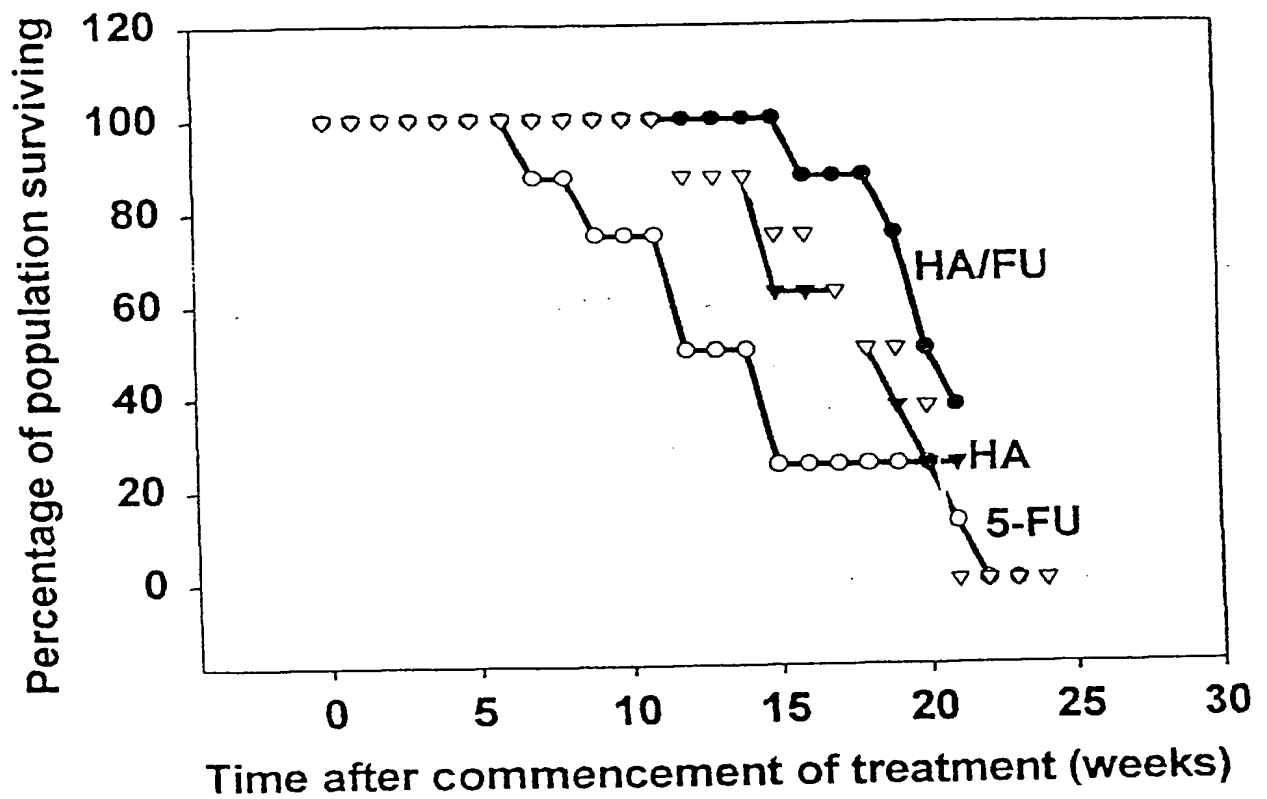


FIGURE 21

Effect of CMF/HA therapy on body mass:
6-week study

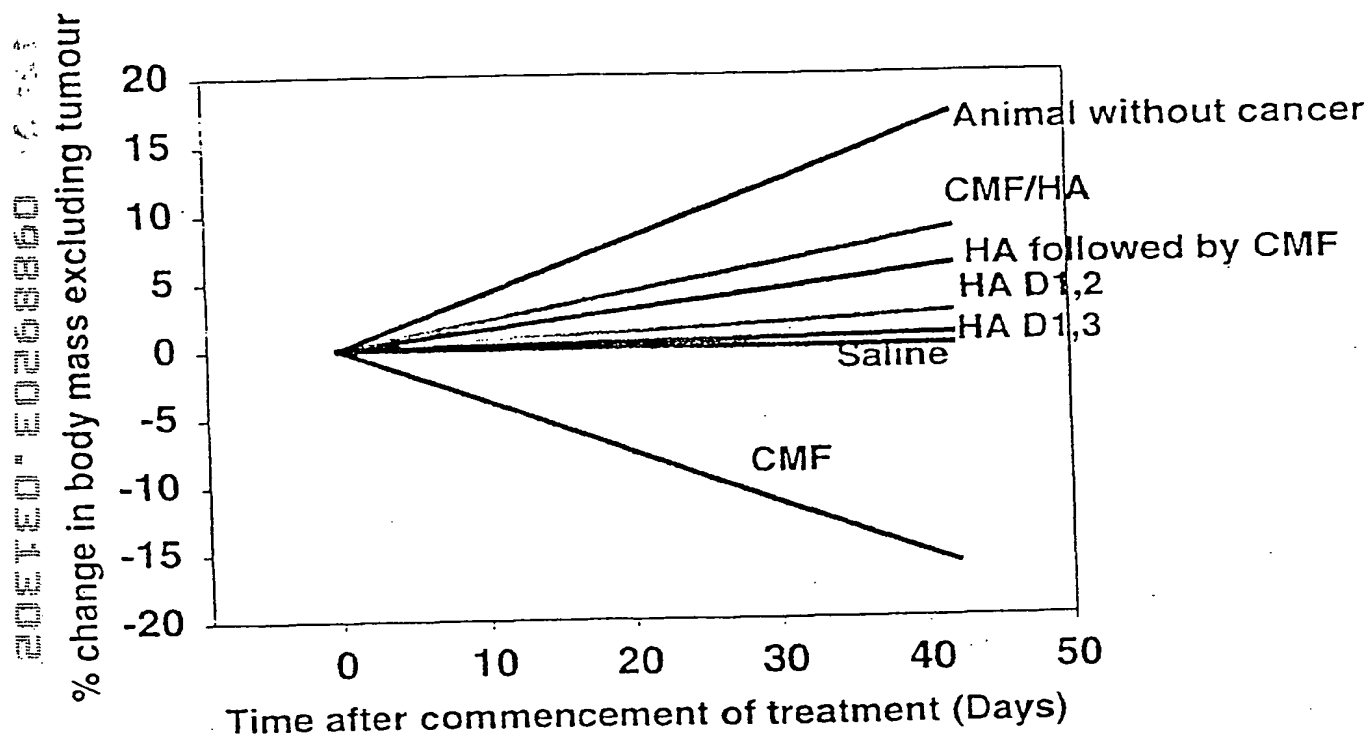


FIGURE 22

Effect of CMF/HA adjuvant therapy on primary tumour mass

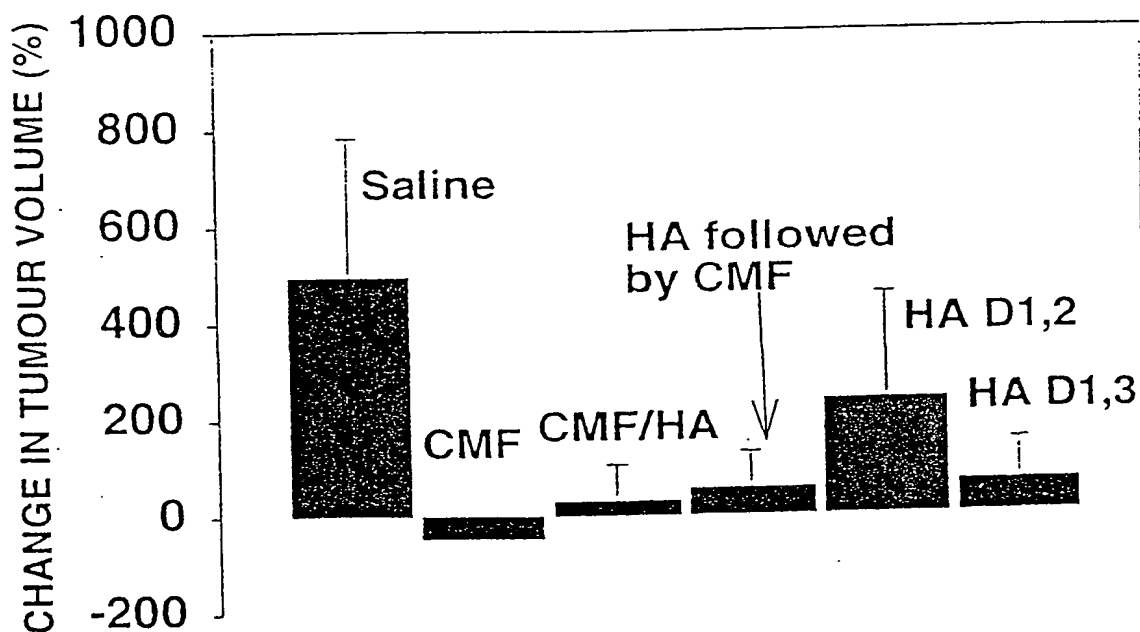


FIGURE 23

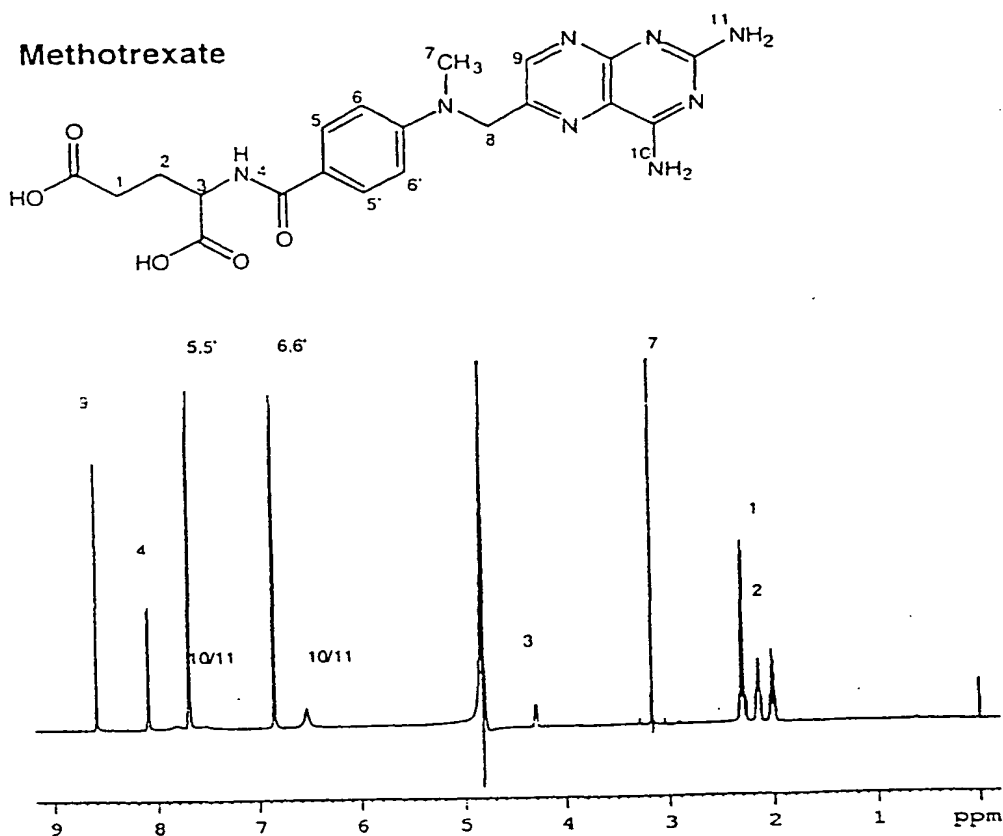


FIGURE 24

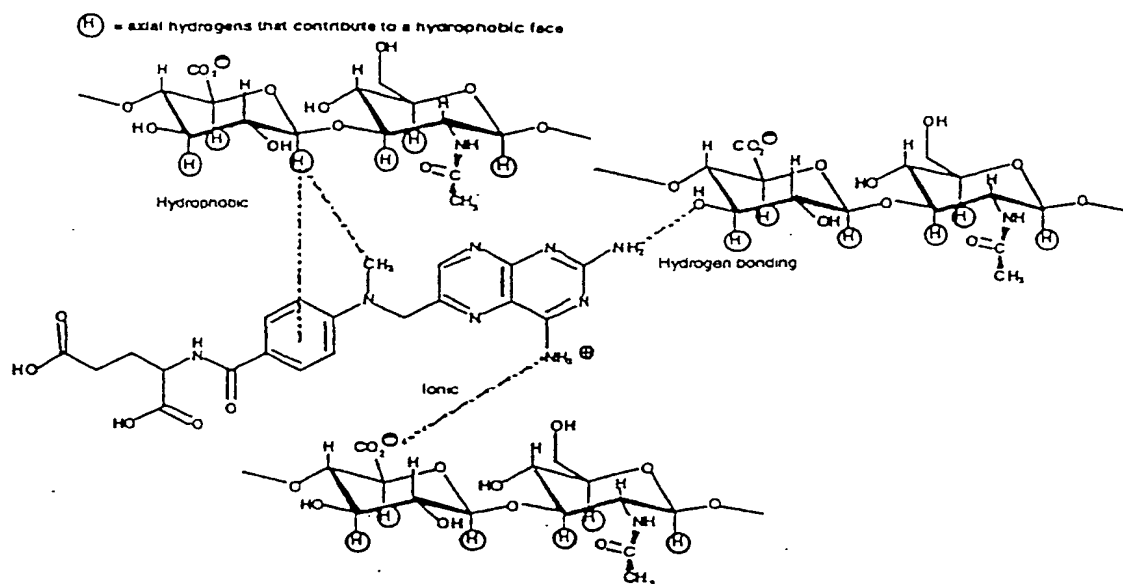


FIGURE 25

09/889203-031302

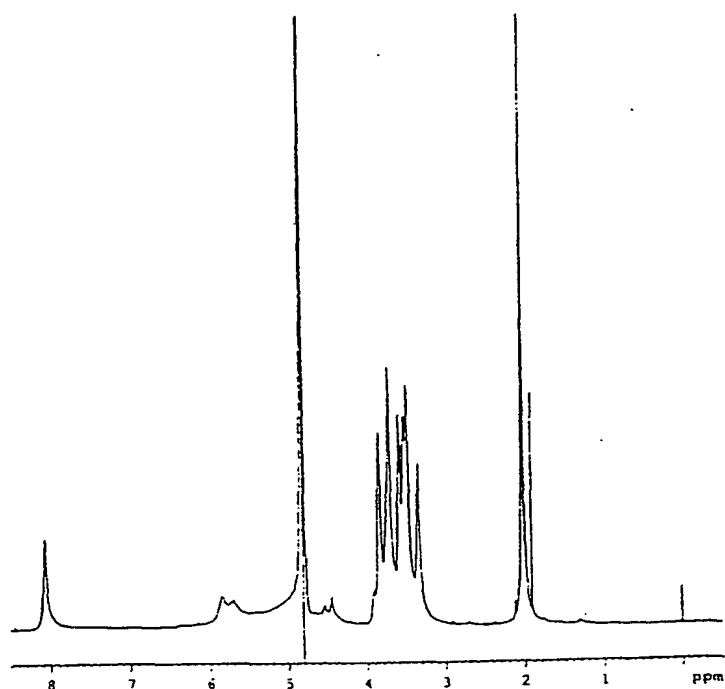


FIGURE 26

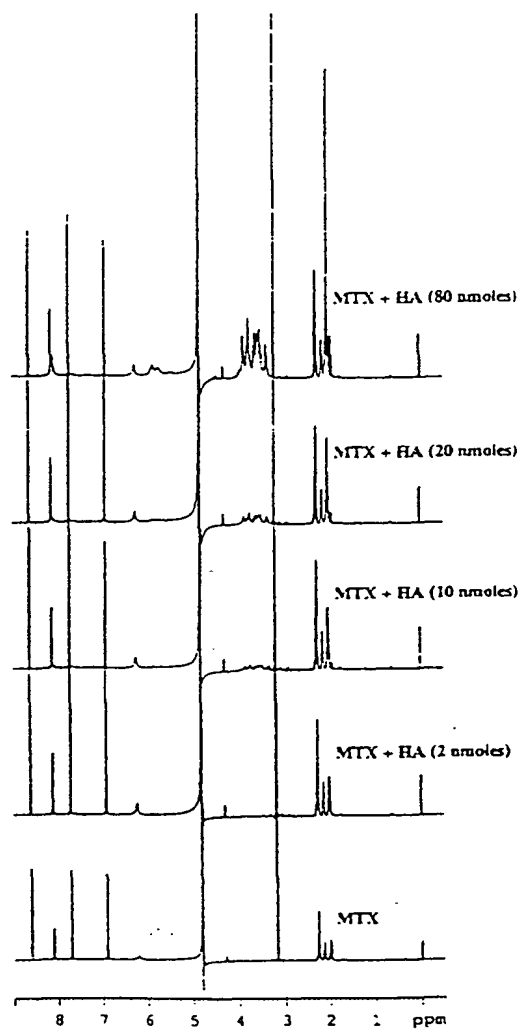


FIGURE 27

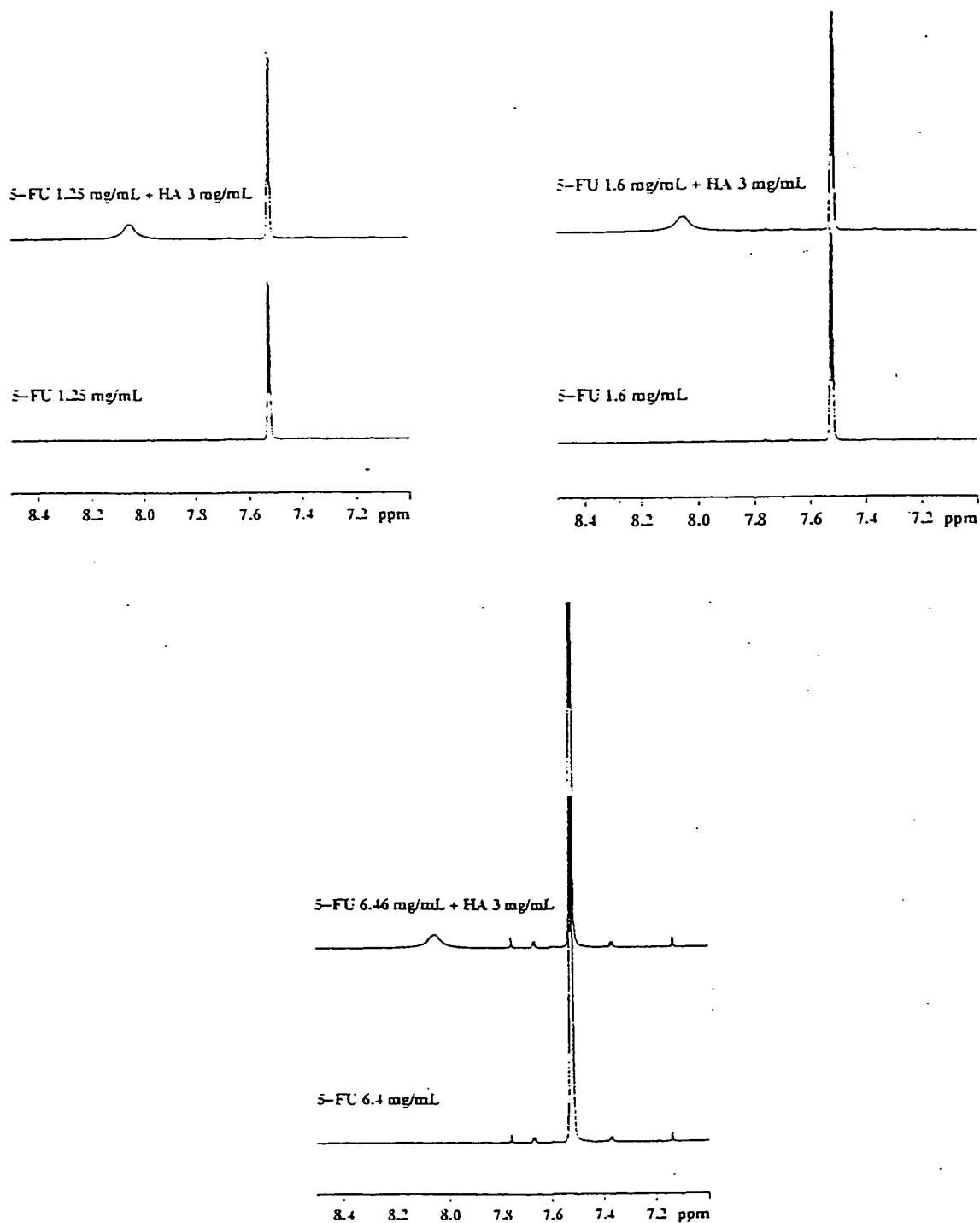


FIGURE 28